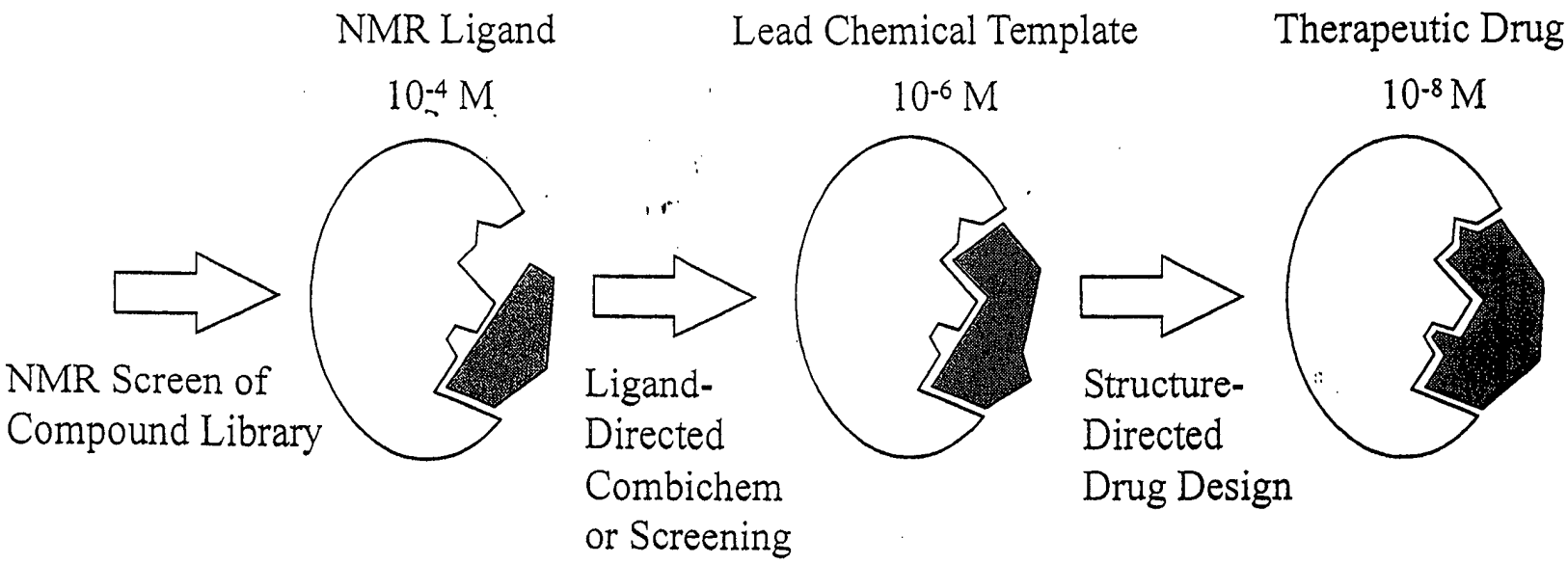
Figure 1



Title: METHODS FOR CREATING A COMPOUND LIBRARY
Applicant(s): Stockman et al.
Serial No.: Unassigned
Express Mail No.: EL88...1489US
Filed: Herewith
Docket: 6283NC
Sheet 1 of 17

HA BCUT

CHRG BCUT

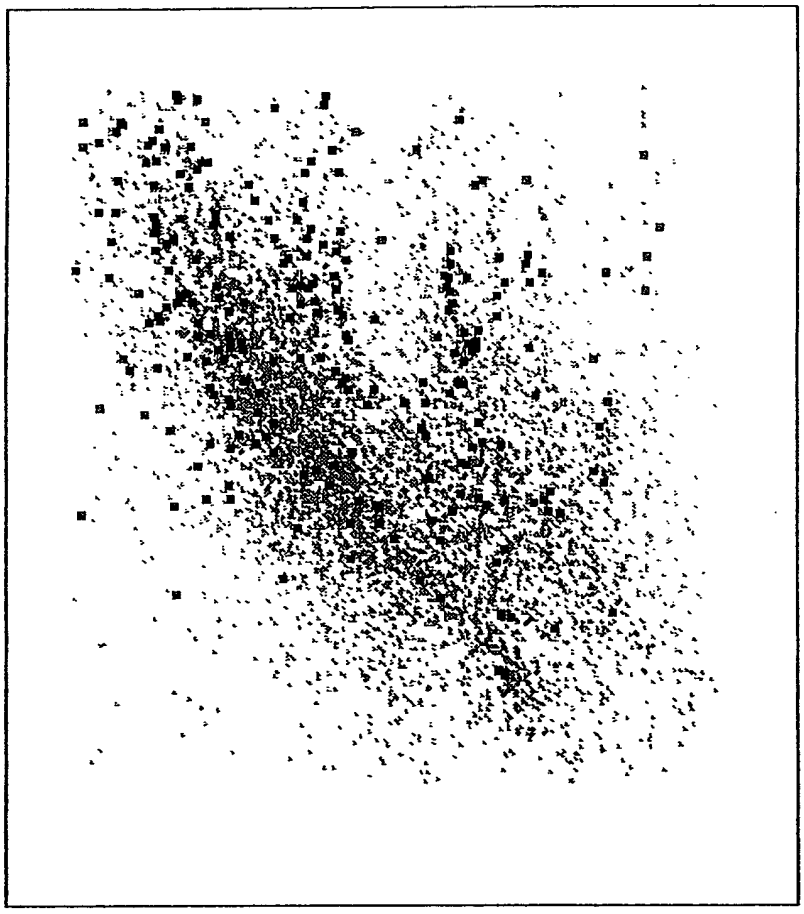
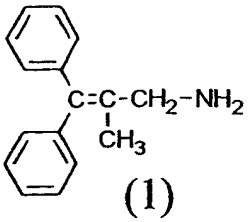
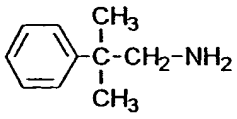


Figure 2

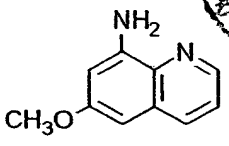
Figure 3


C=C-CH2-NH2
CH3

(1)


CH3
C-CH2-NH2
CH3

(2)


NH2
N
CH3O

(3)

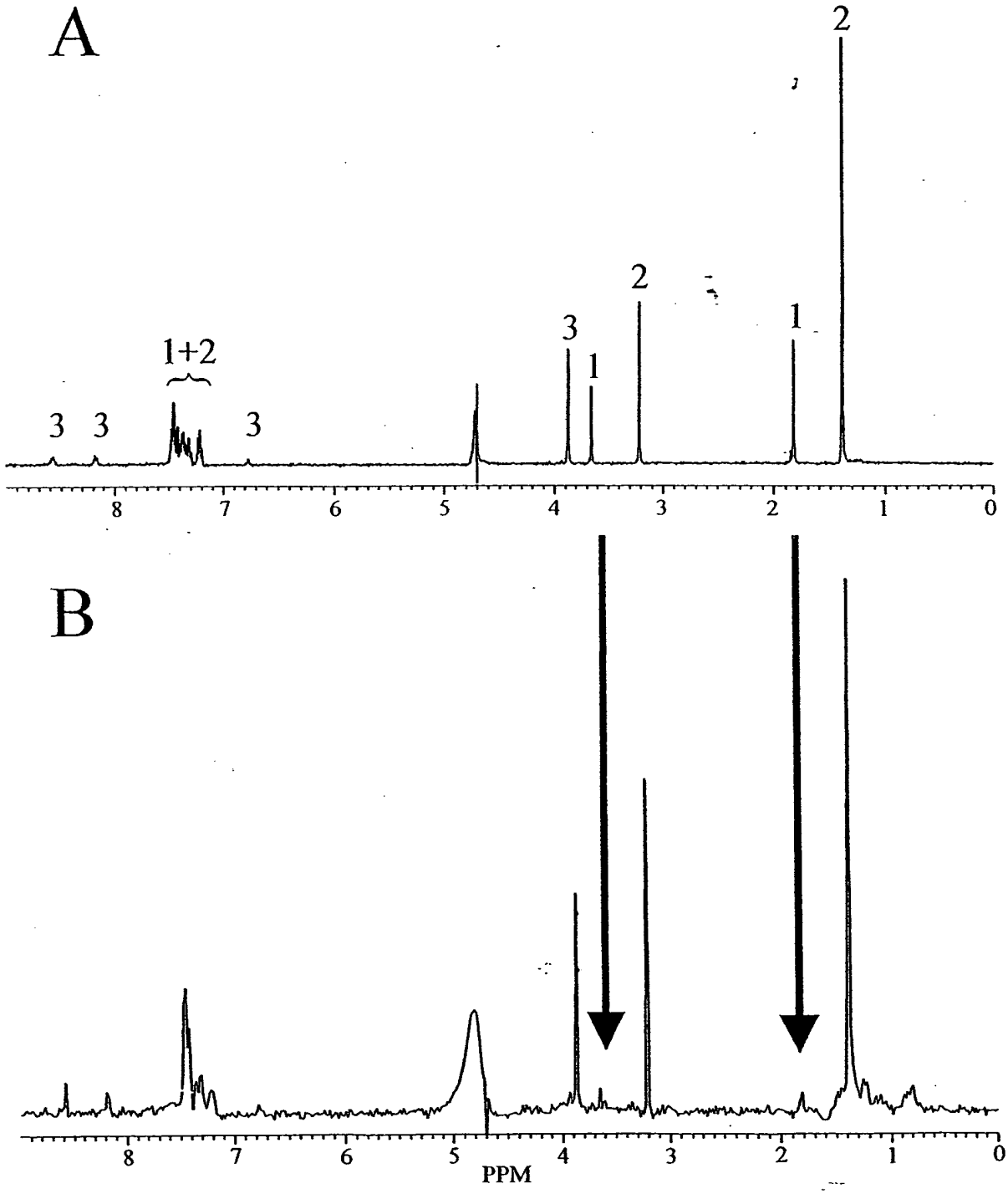
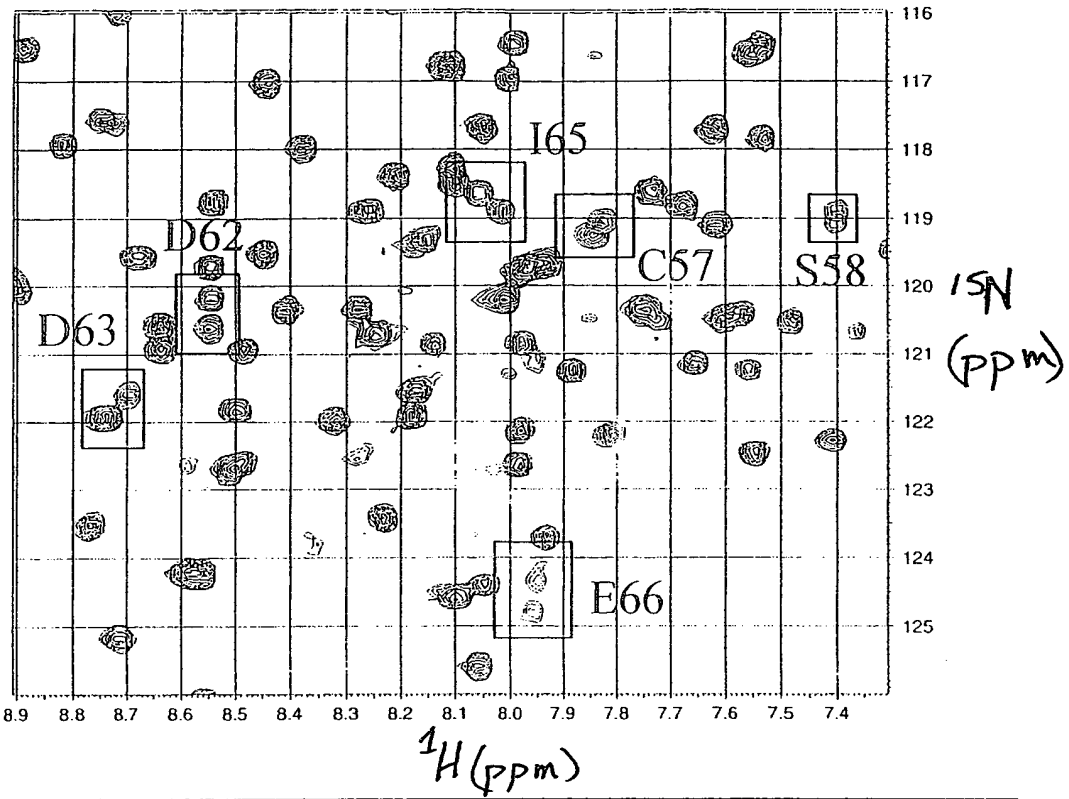

A
1+2
3
3
3
3
1
2
1
2
8
7
6
5
4
3
2
1
0
B
8
7
6
5
4
3
2
1
0
PPM

Figure 4

A


I65
D62
C57
S58
D63
E66
116
117
118
119
120
121
122
123
124
125
15N (ppm)
8.9 8.8 8.7 8.6 8.5 8.4 8.3 8.2 8.1 8.0 7.9 7.8 7.7 7.6 7.5 7.4
1H (ppm)

B

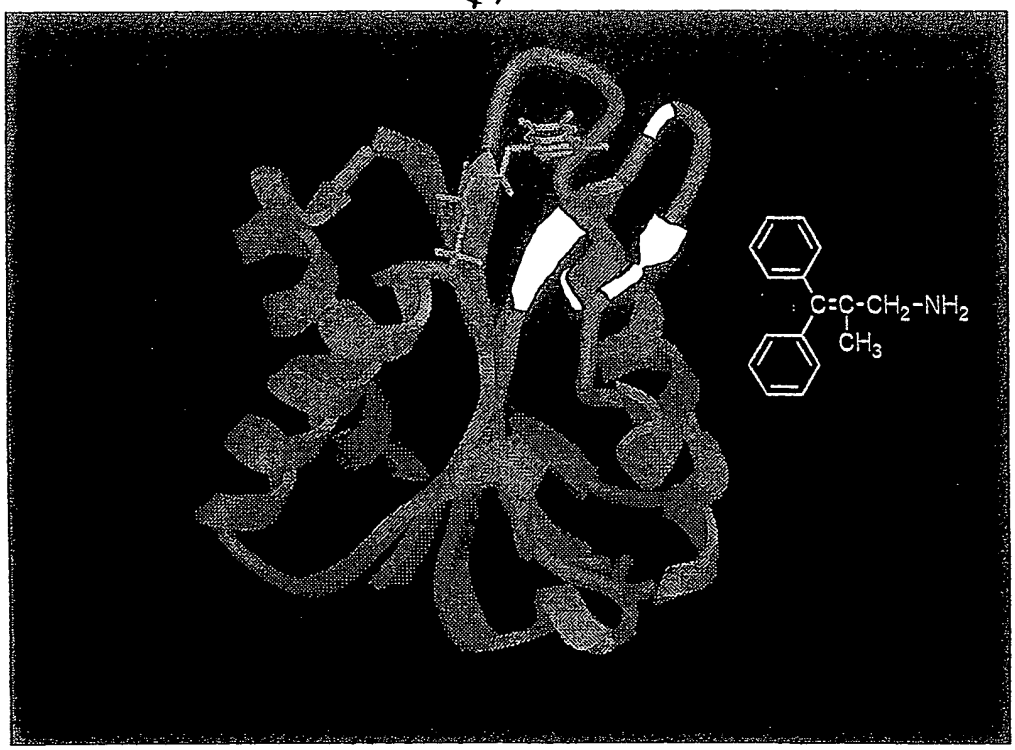
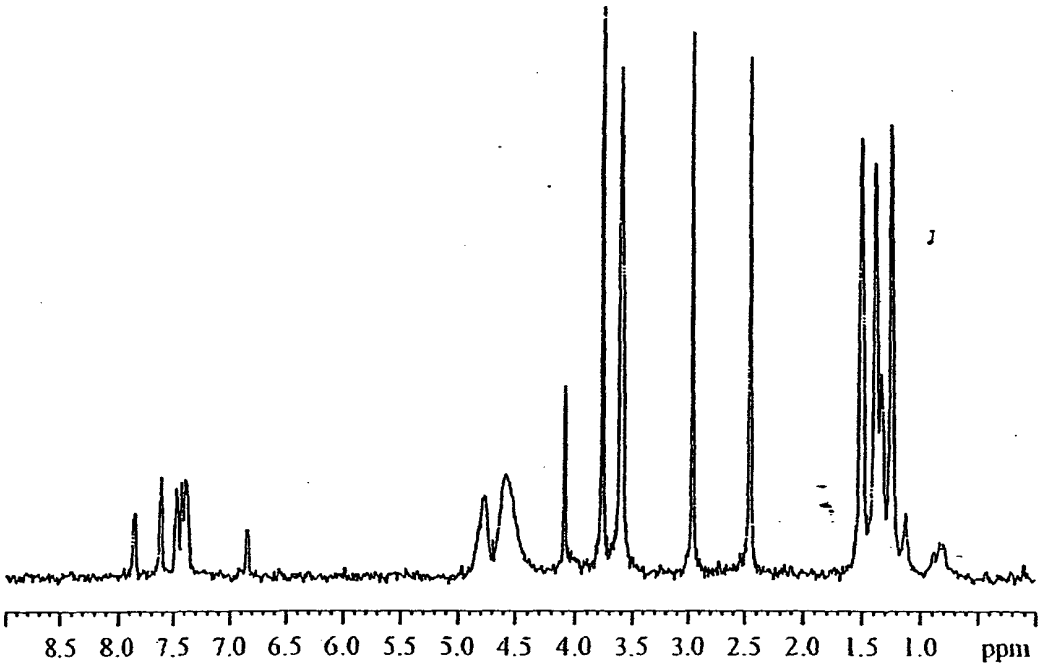

C=C-CH2-NH2
CH3

Figure 5

A


8.5 8.0 7.5 7.0 6.5 6.0 5.5 5.0 4.5 4.0 3.5 3.0 2.5 2.0 1.5 1.0 ppm

B

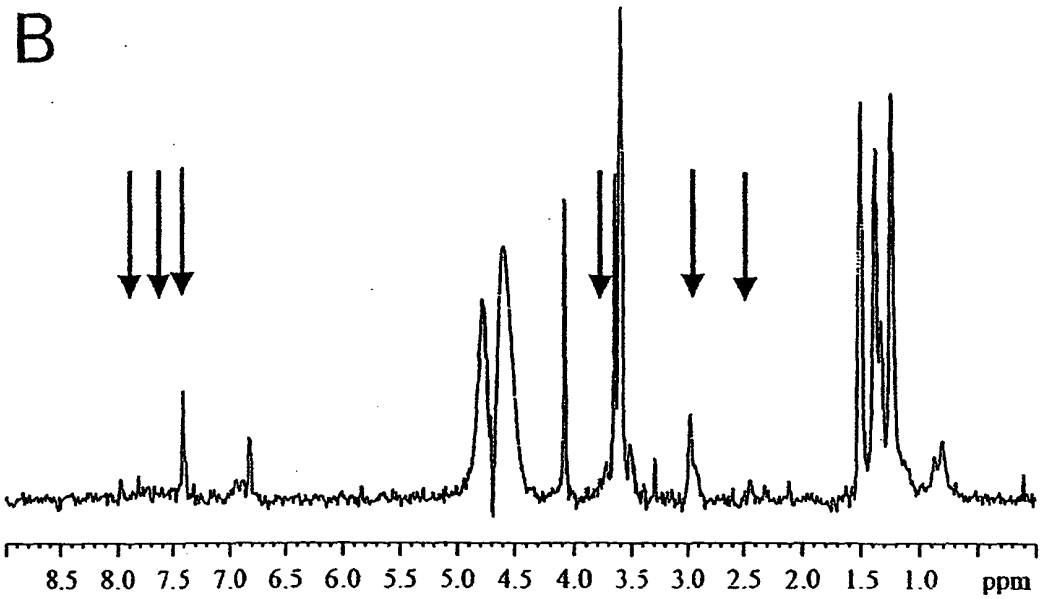

8.5 8.0 7.5 7.0 6.5 6.0 5.5 5.0 4.5 4.0 3.5 3.0 2.5 2.0 1.5 1.0 ppm

Figure 6

Ligand A NMR $K_d \sim 1.4 \times 10^{-4}$ M
Ligand A $IC_{50} \sim 9.0 \times 10^{-5}$ M
Ligand B $IC_{50} = 1.8 \times 10^{-5}$ M
Ligand C $IC_{50} = 1.8 \times 10^{-5}$ M
Ligand D $IC_{50} = 1.2 \times 10^{-5}$ M
Ligand E $IC_{50} = 3.2 \times 10^{-6}$ M

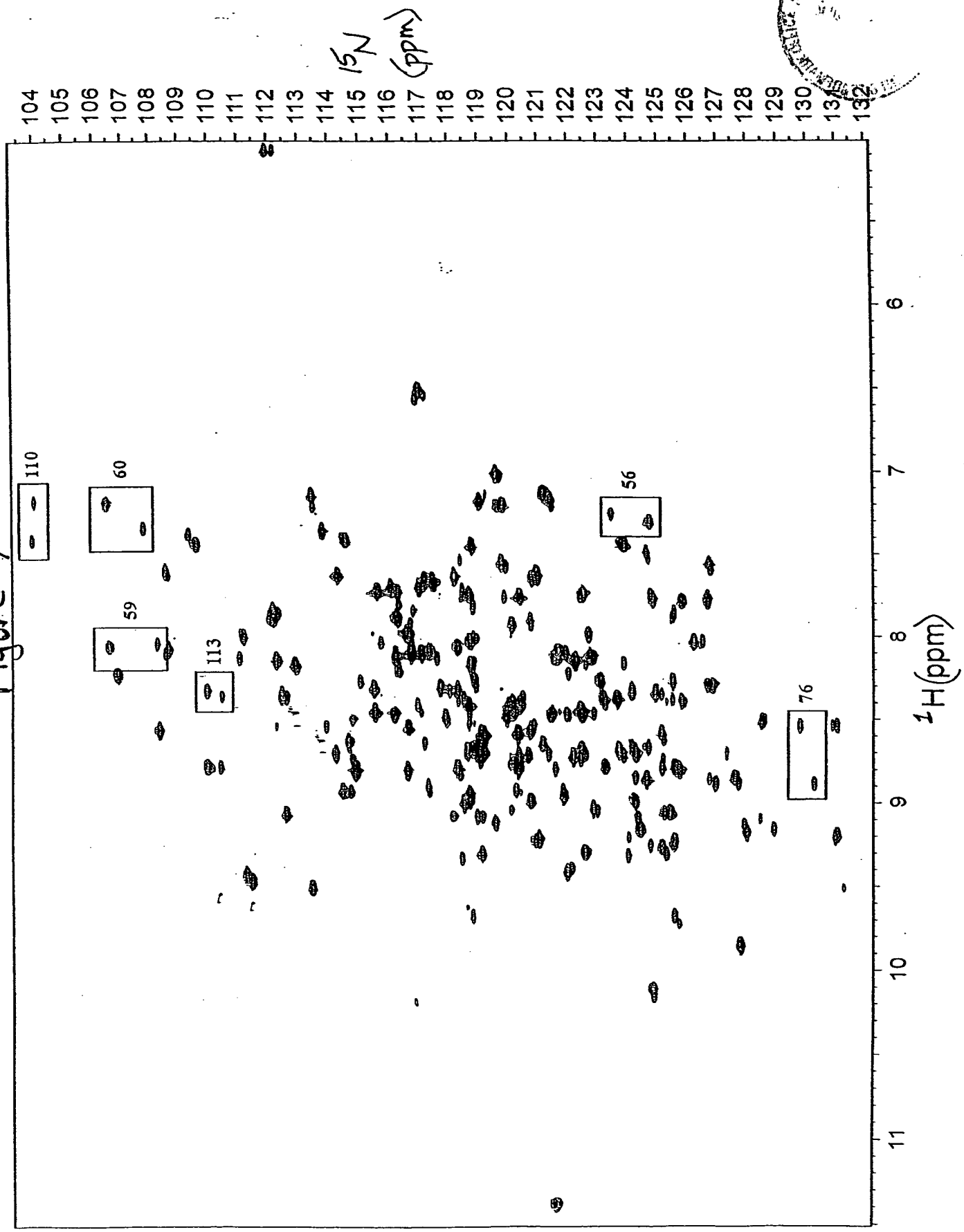

15N (ppm)
104 105 106 107 108 109 110 111 112 113 114 115 116 117 118 119 120 121 122 123 124 125 126 127 128 129 130 131 132
1H (ppm)
6 7 8 9 10 11
110
60
59
113
56
76

Figure 7

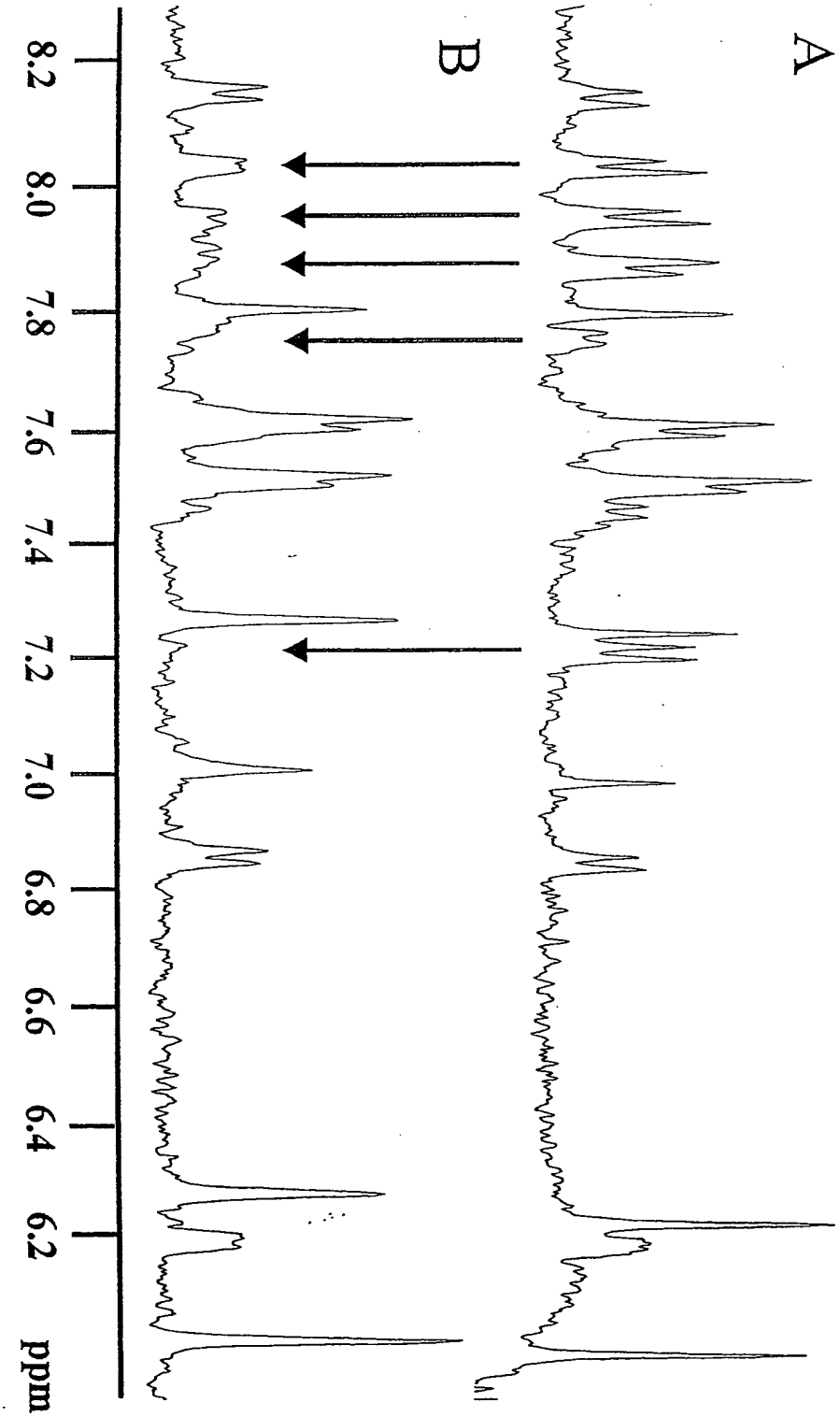


Figure 8

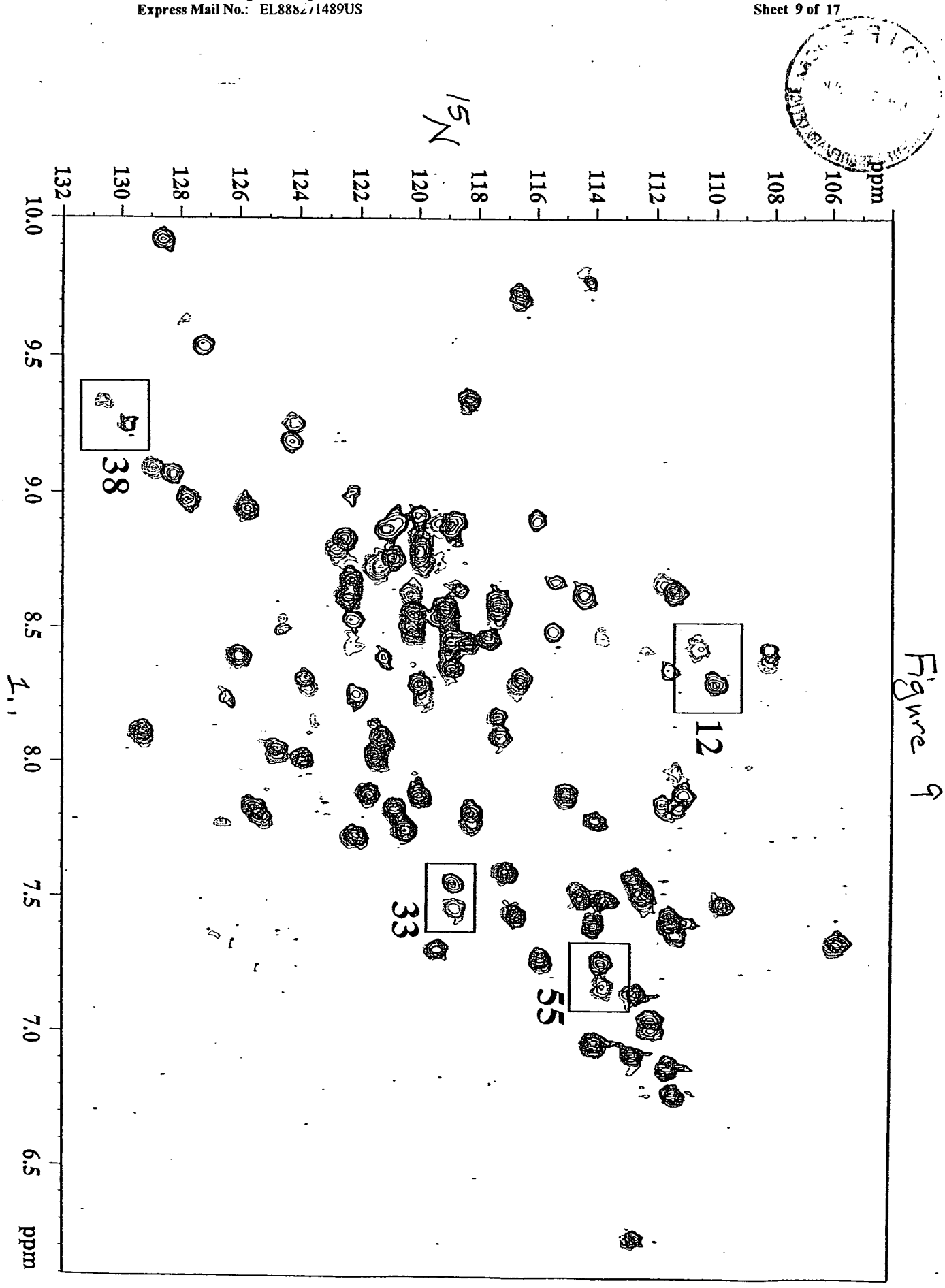

15N
ppm
132
130
128
126
124
122
120
118
116
114
112
110
108
106
10.0
9.5
9.0
8.5
8.0
7.5
7.0
6.5
ppm
38
12
33
55

Figure 9

Figure 10.

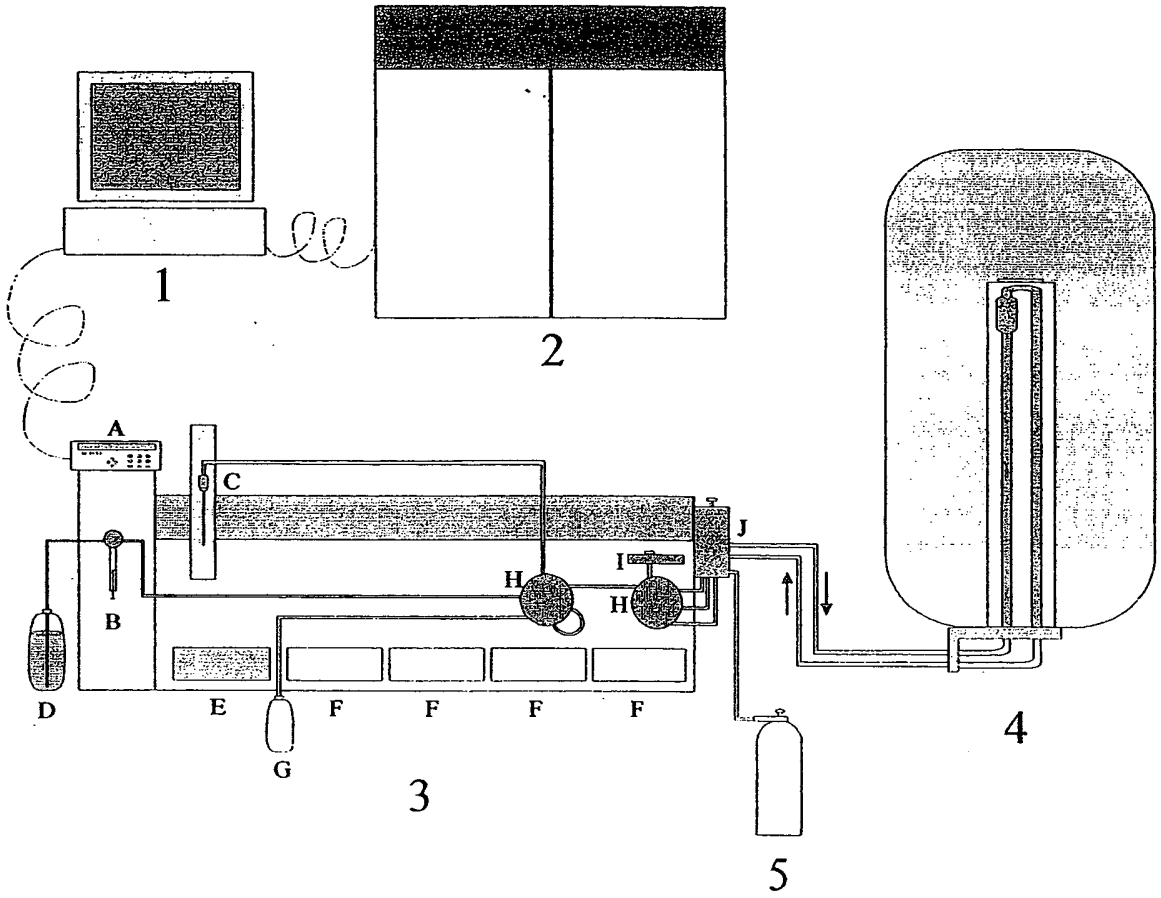


Figure 11.

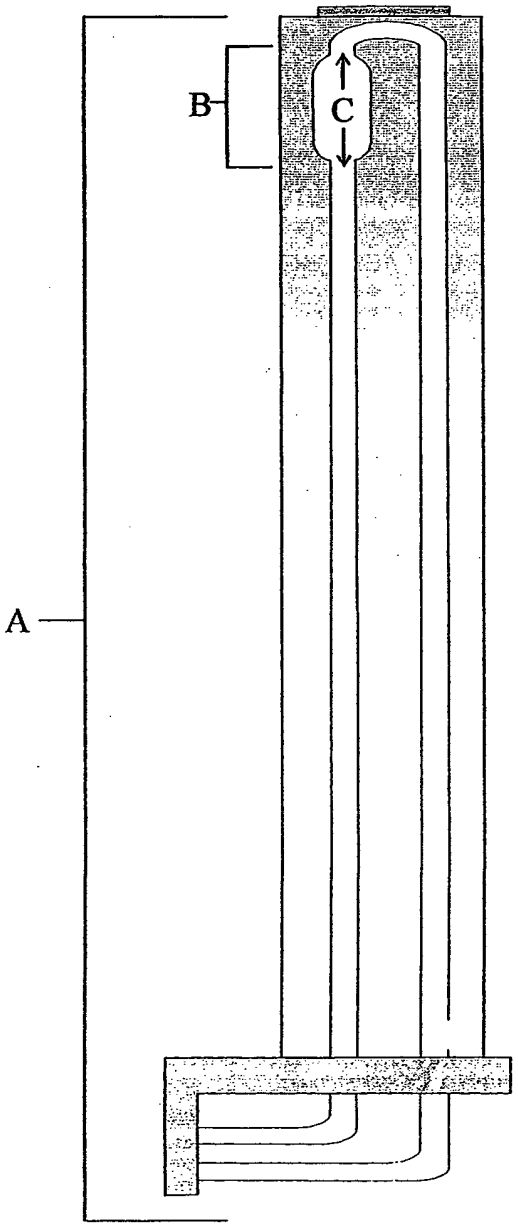


Figure 12.

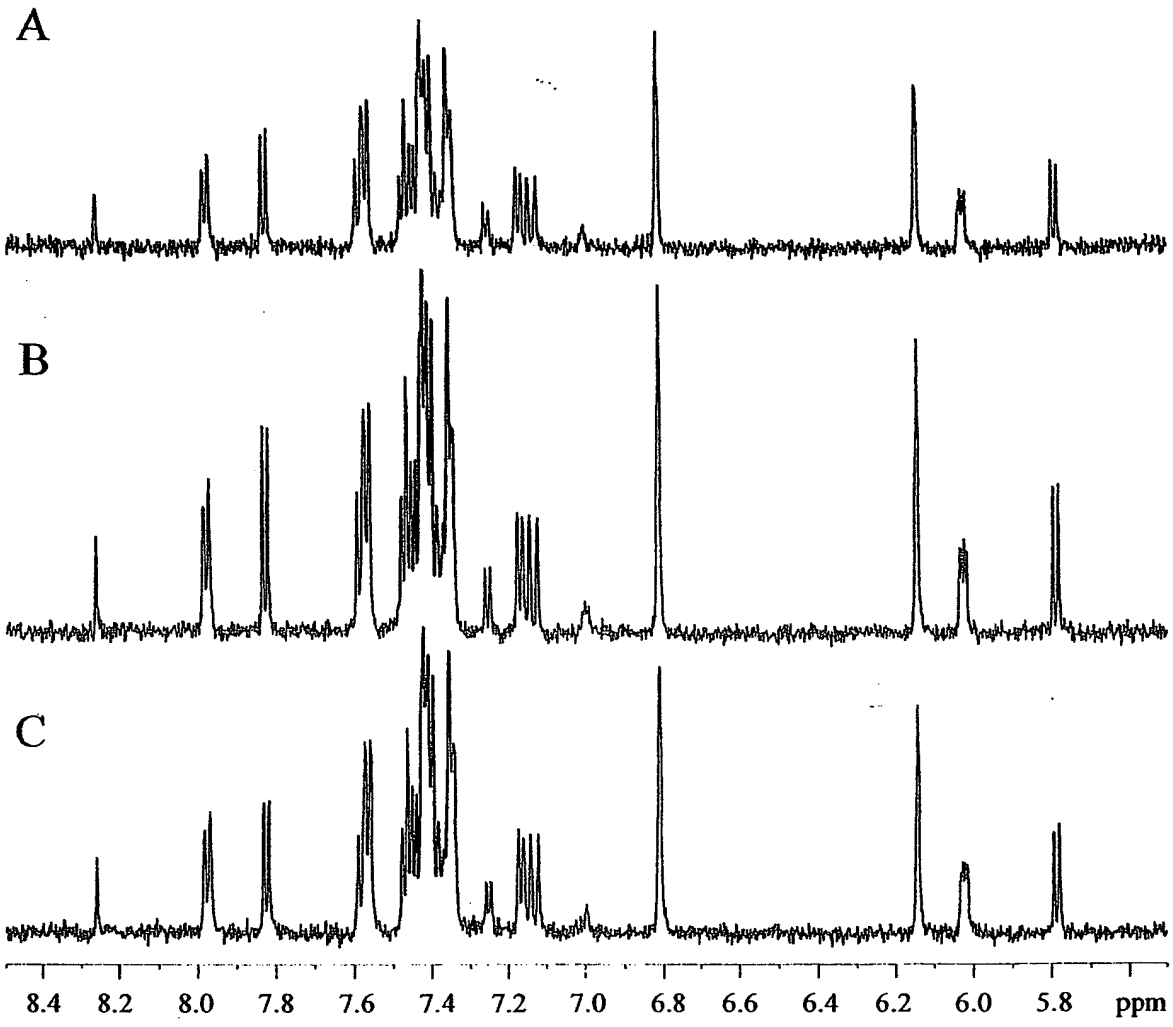


Figure 13.

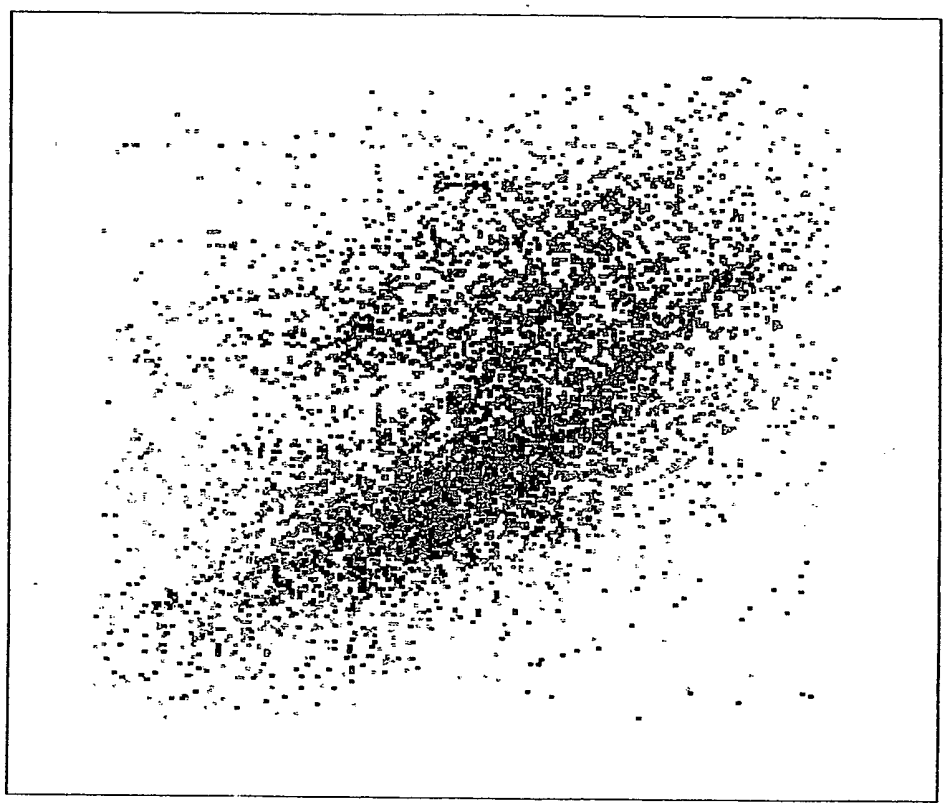


Figure 14.

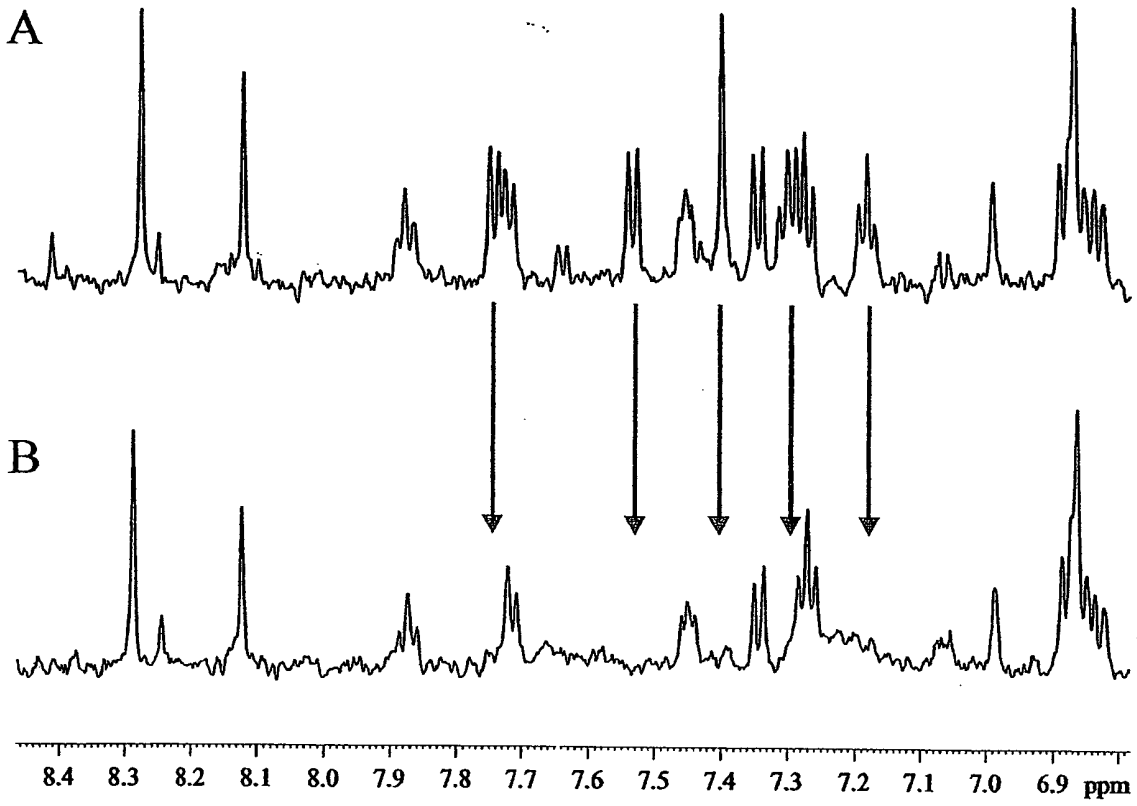


Figure 16

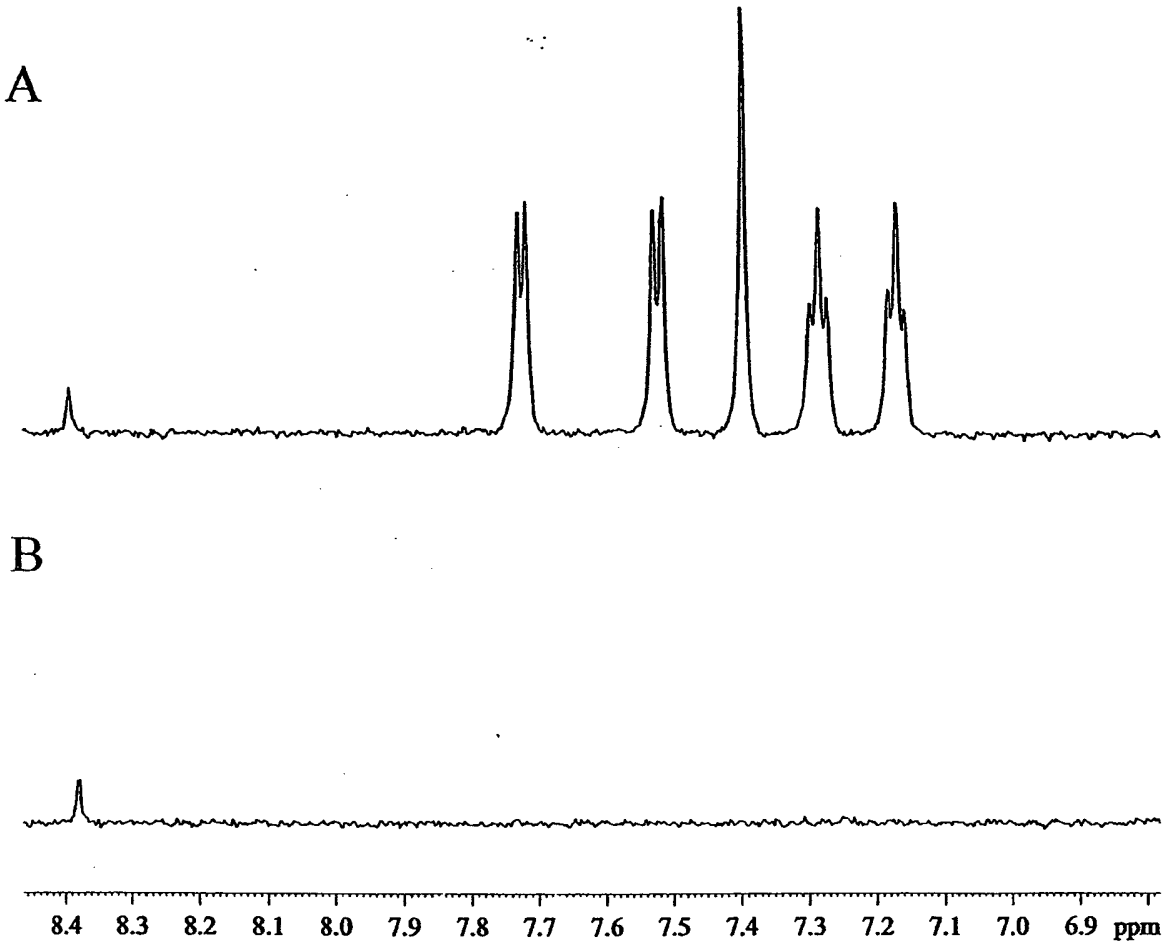
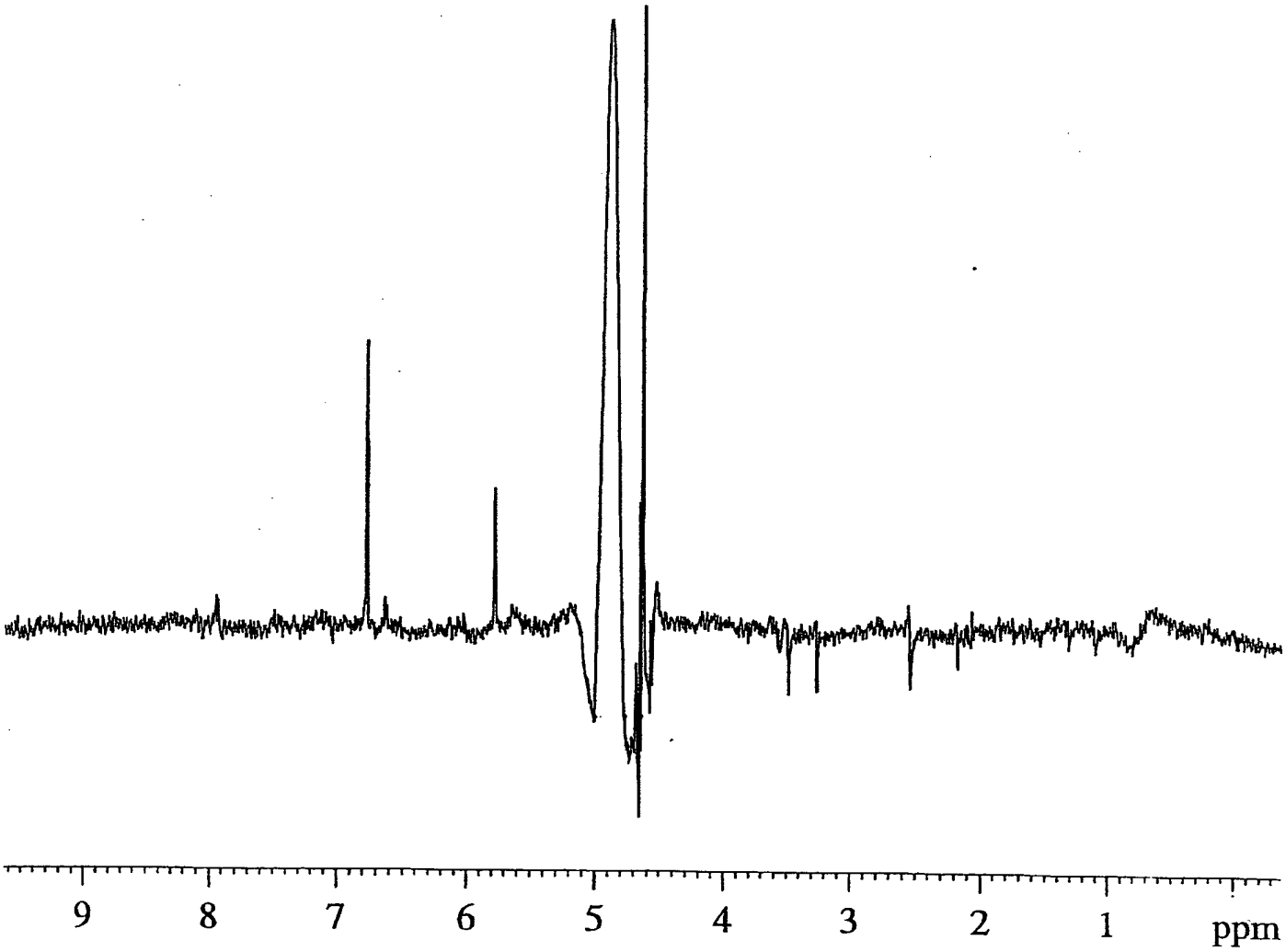


Figure 16



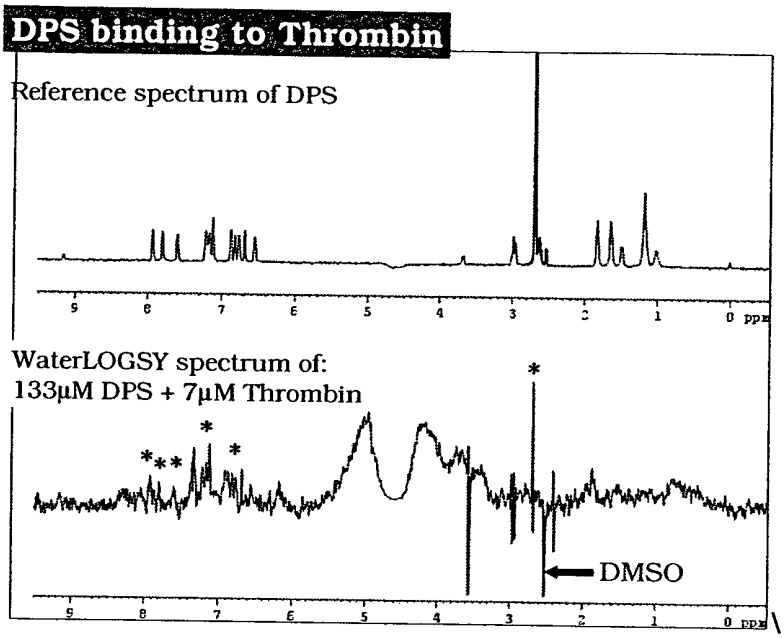

DPS binding to Thrombin
Reference spectrum of DPS
9 8 7 6 5 4 3 2 1 0 ppm
WaterLOGSY spectrum of:
133μM DPS + 7μM Thrombin
DMSO
9 8 7 6 5 4 3 2 1 0 ppm

Figure 17